

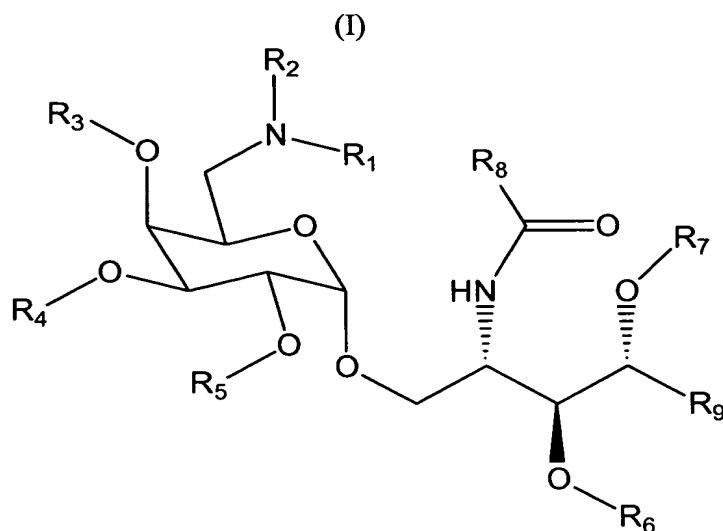
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JC17 Rec'd PCT/PTO 20 SEP 2005

Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

Listing of Claims:

1. (Original) A compound of Formula (I):



wherein,

R₁ is:

(i) hydrogen; or

(ii) -SO₂R₁₀,

wherein R₁₀ is:

halo; hydroxy; OR₁₁; OR₁₂; amino; NHR₁₁; N(R₁₁)₂; NHR₁₂; N(R₁₂)₂; aralkylamino;

or

C₁-C₁₂ alkyl optionally substituted with halo, hydroxy, oxo, nitro, OR₁₁, OR₁₂,

acyloxy, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto,

thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate,

phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl containing 0-3 R₁₄, or heteroaryl containing 0-3 R₁₄; or C₃-C₁₀ cycloalkyl, C₃-C₁₀ heterocyclyl, C₅-C₁₀ cycloalkenyl, or C₅-C₁₀ heterocycloalkenyl optionally substituted with one or more halo, hydroxy, oxo, OR₁₁, OR₁₂, acyloxy, nitro, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto, thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate, phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, alkyl, haloalkyl, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl heteroaryl containing 0-3 R₁₄, or C₆-C₂₀ heteroaryl containing 0-3 R₁₄; or C₂-C₆ alkenyl, C₂-C₆ alkynyl, aryl, or heteroaryl optionally substituted with one or more halo, hydroxy, OR₁₁, OR₁₂, acyloxy, nitro, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto, thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate, phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, alkyl, haloalkyl, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl containing 0-3 R₁₄, or C₆-C₂₀ heteroaryl containing 0-3 R₁₄; or

(iii) -C(O)R₁₀, wherein R₁₀ is defined as above; or

(iv) -C(R₁₀)₂(R₁₅), wherein R₁₀ is defined as above; R₁₅ is hydrogen, R₁₀, or R₁₅ and R₂ taken together forms a double bond between the carbon and nitrogen atoms to which they are attached; or

(v) R₁ and R₂ taken together forms a heterocyclyl of 3-10 ring atoms optionally substituted with R₁₀;

R₂ is hydrogen, or R₂ and R₁₅ taken together forms a double bond between the carbon and nitrogen atoms to which they are attached, or R₂ and R₁ taken together forms a heterocyclyl of 3-10 ring atoms optionally substituted with R₁₀;

R₃, R₄, R₅, R₆, and R₇ are each independently hydrogen, C₁-C₆ alkyl, C₆-C₁₂ aralkyl, or C₁-C₆ acyl;

R₈ is $-(CH_2)_xCH_3$;

R₉ is a linear or branched C₃-C₁₀₀ alkyl;

R₁₁ is C₁-C₂₀ alkyl optionally substituted with halo, hydroxy, alkoxy, amino, alkylamino, dialkylamino, sulfate, or phosphate;

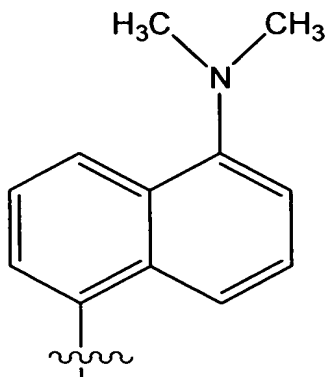
R₁₂ is aryl optionally substituted with halo, haloalkyl, hydroxy, alkoxy, nitro, amino, alkylamino, dialkylamino, sulfate, or phosphate;

Each R₁₃ is independently halo, haloalkyl, hydroxy, alkoxy, oxo, amino, alkylamino, dialkylamino, sulfate, or phosphate;

Each R₁₄ is independently halo, haloalkyl, hydroxy, alkoxy, nitro, amino, alkylamino, dialkylamino, sulfate, or phosphate; and

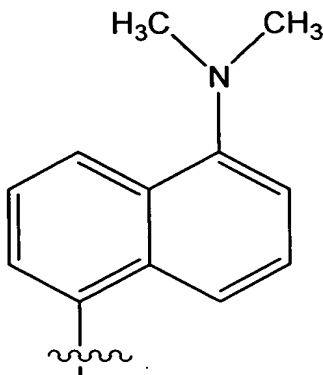
x is 1-100.

2. (Original) The compound of claim 1 wherein x is 24 and R₉ is *n*-tetradecyl.
3. (Original) The compound of claim 2 wherein R₁ is SO₂R₁₀.
4. (Currently Amended) The compound of claim 3 wherein R₁₀ is aryl substituted with N(R₁₁)₂[;].
5. (Currently Amended) The compound of claim 4 wherein R₁₀ is:



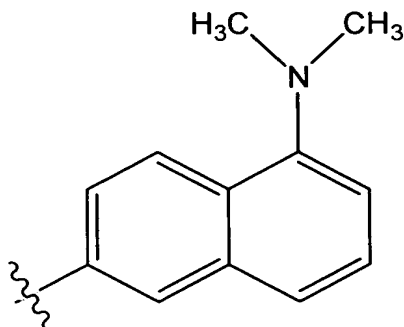
6. (Original) The compound of claim 2 wherein R₁ is C(O)R₁₀.
7. (Currently Amended) The compound of claim 6 wherein R₁₀ is C₁-C₆ alkyl substituted with halo, hydroxy, oxo, nitro, OR₁₁, OR₁₂, acyloxy, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto, thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate, phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl containing 0-3 R₁₄, or C₆-C₂₀ heteroaryl containing 0-3 R₁₄[[i]]].
8. (Original) The compound of claim 7 wherein R₁₀ is C₁-C₆ alkyl substituted with NHSO₂R₁₂.

9. (Currently Amended) The compound of claim 8 wherein R_{12} is:



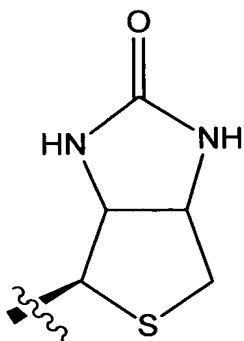
10. (Original) The compound of claim 7, wherein R_{10} is alkyl substituted with $C(O)R_{12}$.

11. (Currently Amended) The compound of claim 10 wherein R_{12} is:

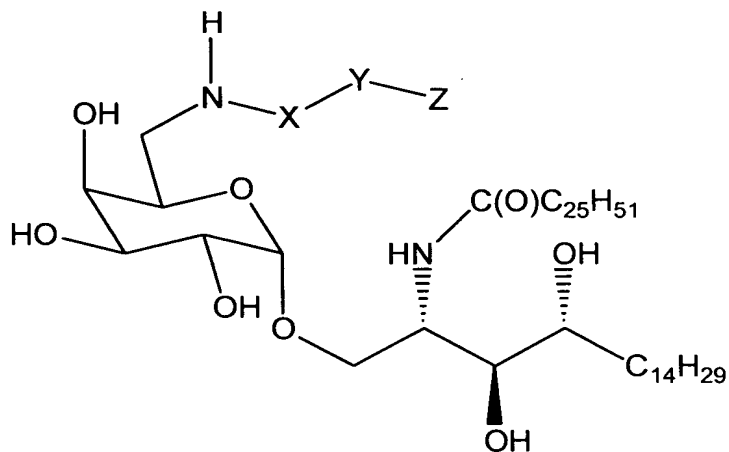


12. (Original) The compound of claim 7 wherein R_{10} is alkyl is substituted with C_5 - C_{10} heterocyclyl containing 0-3 R_{13} .

13. (Currently Amended) The compound of claim 12 wherein the heterocyclyl is:



14. (Original) A probe for observing glycolipid association with CD1d and NKT cell receptors during NKT cell stimulation comprising a compound of Formula (II):



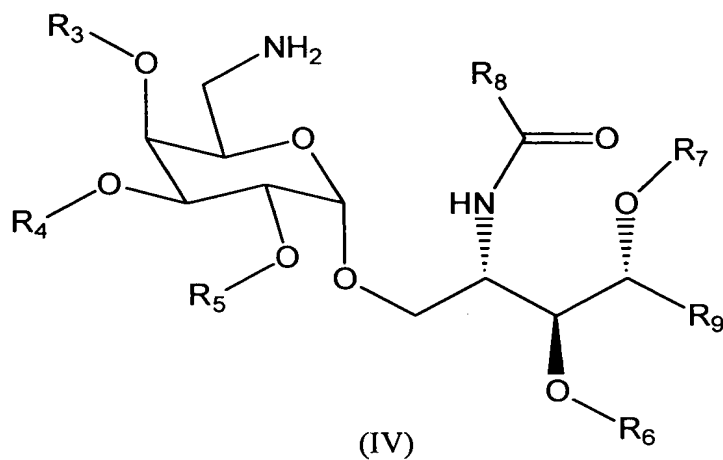
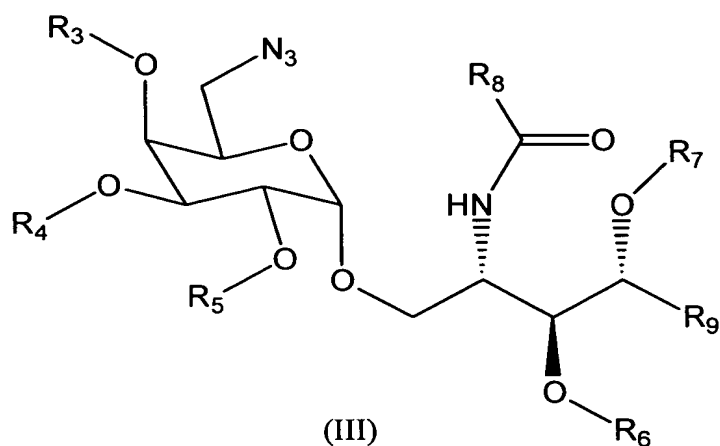
wherein:

X is $-\text{SO}_2-$, $-\text{C}(\text{O})-$, or absent;

Y is a linker group; and

Z is a reporter group.

15. (Original) A method of quantifying glycolipid association with CD1d and NKT cell receptors during NKT cell stimulation comprising: (i) contacting a compound of Formula (II) with a CD1d protein; (ii) allowing the compound to associate with the CD1d protein; (iii) measuring fluorescence emitted by the compound during steps (i) and (ii) to provide one or more pre-NKT cell contact fluorescence measurements; (iv) contacting the compound and CD1d protein with an NKT cell line; (v) measuring fluorescence emitted by the compound during step (iv) to provide one or more NKT cell contact fluorescence measurements.
16. (Original) The method of claim 15 wherein step (v) is repeated over time.
17. (Original) The method of claim 15 further comprising the step of comparing the fluorescence measurements in steps (iii) and (v).
18. (Original) A method of stimulating NKT cells comprising contacting an NKT cell with a compound of Formula (I) and a CD1 protein.
19. (Original) The method of claim 18 wherein the protein is CD1d.
20. (Original) A method of stimulating the immune system of a subject in need of such stimulation, the method comprising administering a compound of Formula (I) to the subject.
21. (Original) A method of treating an autoimmune disease in a subject in need of such treatment, the method comprising administering an effective amount of a compound of Formula (I).
22. (Currently Amended) The method of claim ~~20~~ or 21 wherein the subject is a mammal.
23. (Original) The method of claim 22 wherein the subject is a human.
24. (Original) A method of making a compound of Formula (I) comprising: (i) converting a compound of Formula (III) to a compound of Formula (IV):



and (ii) contacting a compound of Formula (IV) with R₁-LG to afford a compound of Formula (I), wherein:

R₁ is:

(i) -SO₂R₁₀,

wherein R₁₀ is:

halo; hydroxy; OR₁₁; OR₁₂; amino; NHR₁₁; N(R₁₁)₂; NHR₁₂; N(R₁₂)₂; aralkylamino;

or

C₁-C₁₂ alkyl optionally substituted with halo, hydroxy, oxo, nitro, OR₁₁, OR₁₂, acyloxy, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto, thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate, phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl containing 0-3 R₁₄, or C₆-C₂₀ heteroaryl containing 0-3 R₁₄; or C₃-C₁₀ cycloalkyl, C₃-C₁₀ heterocyclyl, C₅-C₁₀ cycloalkenyl, or C₅-C₁₀ heterocycloalkenyl optionally substituted with one or more halo, hydroxy, oxo, OR₁₁, OR₁₂, acyloxy, nitro, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto, thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate, phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, alkyl, haloalkyl, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl containing 0-3 R₁₄, or C₆-C₂₀ heteroaryl containing 0-3 R₁₄; or C₂-C₆ alkenyl, C₂-C₆ alkynyl, aryl, or heteroaryl optionally substituted with one or more halo, hydroxy, OR₁₁, OR₁₂, acyloxy, nitro, amino, NHR₁₁, N(R₁₁)₂, NHR₁₂, N(R₁₂)₂, aralkylamino, mercapto, thioalkoxy, S(O)R₁₁, S(O)R₁₂, SO₂R₁₁, SO₂R₁₂, NHSO₂R₁₁, NHSO₂R₁₂, sulfate, phosphate, cyano, carboxyl, C(O)R₁₁, C(O)R₁₂, C(O)OR₁₁, C(O)NH₂, C(O)NHR₁₁, C(O)N(R₁₁)₂, alkyl, haloalkyl, C₃-C₁₀ cycloalkyl containing 0-3 R₁₃, C₃-C₁₀ heterocyclyl containing 0-3 R₁₃, C₂-C₆ alkenyl, C₂-C₆ alkynyl, C₅-C₁₀ cycloalkenyl, C₅-C₁₀ heterocycloalkenyl, C₆-C₂₀ aryl containing 0-3 R₁₄, or C₆-C₂₀ heteroaryl containing 0-3 R₁₄; or

(ii) -C(O)R₁₀, wherein R₁₀ is defined as above; or

(iii) -C(R₁₀)₂(R₁₅), wherein R₁₀ is defined as above; R₁₅ is hydrogen, R₁₀, or R₁₅ and

R₂ taken together forms a double bond between the carbon and nitrogen atoms to which they are attached; or

R₃, R₄, R₅, R₆, and R₇ are each independently hydrogen, C₁-C₆ alkyl, C₆-C₁₂ aralkyl, or C₁-C₆ acyl;

R_8 is $-(CH_2)_xCH_3$;

R_9 is a linear or branched C_3 - C_{100} alkyl;

R_{11} is C_1 - C_{20} alkyl optionally substituted with halo, hydroxy, alkoxy, amino, alkylamino, dialkylamino, sulfate, or phosphate;

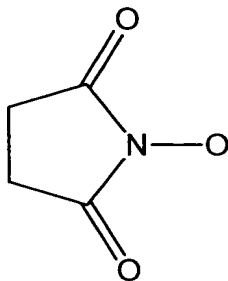
R_{12} is aryl optionally substituted with halo, haloalkyl, hydroxy, alkoxy, nitro, amino, alkylamino, dialkylamino, sulfate, or phosphate;

Each R_{13} is independently halo, haloalkyl, hydroxy, alkoxy, oxo, amino, alkylamino, dialkylamino, sulfate, or phosphate;

Each R_{14} is independently halo, haloalkyl, hydroxy, alkoxy, nitro, amino, alkylamino, dialkylamino, sulfate, or phosphate;

x is 1-100;

LG is halo, $-OSO_2R_{16}$, $B(OH)_2$, or



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; and

R₁₆ is alkyl, haloalkyl or aryl optionally substituted with alkyl, halo or nitro.

25. (Currently Amended) A pharmaceutical ~~composition~~ composition comprising a compound of Formula (I) and a pharmaceutically acceptable carrier.